

United States Patent and Trademark Office



UNITED STATES DEPARTMENT OF COMMERCE United States Patent and Trademark Office Address: COMMISSIONER FOR PATENTS P.O. Box 1450 Alexandria, Virginia 22313-1450 www.uspto.gov

| APPLICATION NO. | FILING DATE | FIRST NAMED INVENTOR | ATTORNEY DOCKET NO. | CONFIRMATION NO. | | | | | |
|--------------------------|--------------------|----------------------|------------------------|------------------|--|--|--|--|--|
| 09/993,234 | 11/19/2001 | Avi J. Ashkenazi | P1007P1D1 | 1337 | | | | | |
| 9157 759 | . 10/07/2003 | | EXAM | INER | | | | | |
| GENENTECH | , INC. | NICKOL, | NICKOL, GARY B | | | | | | |
| 1 DNA WAY SOUTH SAN F | RANCISCO, CA 94080 | 0 | . ART UNIT | PAPER NUMBER | | | | | |
| | , | | 1642 | | | | | | |
| | | | DATE MAILED: 10/07/200 | \mathcal{C} | | | | | |

Please find below and/or attached an Office communication concerning this application or proceeding.

| 4 ' | | Applicant(s) | | | | | | | | | | | | |
|--|--|----------------------------------|---|--|--|--|--|--|--|--|--|--|--|--|
| 09/993,234 ASHKENAZI, AVI J. | | | | | | | | | | | | | | |
| | Office Action Summary | Examiner | Art Unit | | | | | | | | | | | |
| | | Gary B. Nickol Ph.D. | 1642 | | | | | | | | | | | |
| | The MAILING DATE of this communication app | ears on the cover sheet with the | correspondence address | | | | | | | | | | | |
| Period fo | | | | | | | | | | | | | | |
| THE I - Exter after - If the - If NO - Failu - Any r earne | after SIX (6) MONTHS from the mailing date of this communication. If the period for reply specified above is less than thirty (30) days, a reply within the statutory minimum of thirty (30) days will be considered timely. If NO period for reply is specified above, the maximum statutory period will apply and will expire SIX (6) MONTHS from the mailing date of this communication. Failure to reply within the set or extended period for reply will, by statute, cause the application to become ABANDONED (35 U.S.C. § 133). Any reply received by the Office later than three months after the mailing date of this communication, even if timely filed, may reduce any earned patent term adjustment. See 37 CFR 1.704(b). | | | | | | | | | | | | | |
| Status | December to access to the A7.1 | | | | | | | | | | | | | |
| 1)[\] | Responsive to communication(s) filed on <u>17 Jan</u> | | | | | | | | | | | | | |
| 2a)∐ | ,— | s action is non-final. | | | | | | | | | | | | |
| 3) | Since this application is in condition for allowa closed in accordance with the practice under <i>l</i> | • | | | | | | | | | | | | |
| Dispositi | on of Claims | | | | | | | | | | | | | |
| 4)🖂 | 4) Claim(s) 34-39 and 46-94 is/are pending in the application. | | | | | | | | | | | | | |
| | 4a) Of the above claim(s) <u>35 and 46-94</u> is/are w | vithdrawn from consideration. | | | | | | | | | | | | |
| 5) | 5) Claim(s) is/are allowed. | | | | | | | | | | | | | |
| 6)⊠ | 6)⊠ Claim(s) <u>34 and 36-39</u> is/are rejected. | | | | | | | | | | | | | |
| 7) | Claim(s) is/are objected to. | | | | | | | | | | | | | |
| 8) | Claim(s) are subject to restriction and/or | election requirement. | | | | | | | | | | | | |
| · · | on Papers | | | | | | | | | | | | | |
| | The specification is objected to by the Examiner | | | | | | | | | | | | | |
| 10)[| The drawing(s) filed on is/are: a)☐ accep | | | | | | | | | | | | | |
| 44) 🗆 - | Applicant may not request that any objection to the | <u> </u> | ` , | | | | | | | | | | | |
| 11) | The proposed drawing correction filed on | | oved by the Examiner. | | | | | | | | | | | |
| 12)□ - | If approved, corrected drawings are required in rep | | | | | | | | | | | | | |
| | The oath or declaration is objected to by the Exa | arriner. | | | | | | | | | | | | |
| _ | Inder 35 U.S.C. §§ 119 and 120 | | | | | | | | | | | | | |
| _ | Acknowledgment is made of a claim for foreign | priority under 35 U.S.C. § 119(| a)-(d) or (t). | | | | | | | | | | | |
| a)[| ☐ All b)☐ Some * c)☐ None of: | | | | | | | | | | | | | |
| | 1. Certified copies of the priority documents | | | | | | | | | | | | | |
| | 2. Certified copies of the priority documents | | | | | | | | | | | | | |
| * S | 3. Copies of the certified copies of the prior application from the International Bur see the attached detailed Office action for a list of the ac | eau (PCT Rule 17.2(a)). | | | | | | | | | | | | |
| 14) 🗌 A | acknowledgment is made of a claim for domestic | priority under 35 U.S.C. § 119 | (e) (to a provisional application). | | | | | | | | | | | |
| |) The translation of the foreign language proving the stranslation of the foreign language proving the stranguage | • • | | | | | | | | | | | | |
| Attachmen | t(s) | | | | | | | | | | | | | |
| 2) 🔲 Notic | e of References Cited (PTO-892) e of Draftsperson's Patent Drawing Review (PTO-948) mation Disclosure Statement(s) (PTO-1449) Paper No(s) <u>5</u> . | 5) Notice of Informal | ry (PTO-413) Paper No(s) Patent Application (PTO-152) | | | | | | | | | | | |

Art Unit: 1642

DETAILED ACTION

The response filed on July 17, 2003 (Paper No. 9) to the restriction requirement of March 26, 2003 has been received. Applicant has elected the species corresponding to nucleic acids encoding amino acids 25-198 of SEQ ID NO:6. Because applicant did not distinctly and specifically point out the supposed errors in the restriction requirement, the election has been treated as an election without traverse (MPEP 818.03(a).

Claims 34-39, and 46-94 are pending.

Claims 35, 46-94 are withdrawn from further consideration by the examiner under 37 CFR 1.142(b), as being drawn to a non-elected invention. (It is noted that applicant believes that pending claims 34, 36-39, and 91-94 are readable on the elected invention—see Paper No. 9, page 2. However, nowhere do claims 91-94 recite nucleic acids encoding amino acids 25-198. Thus, claims 91-94 are withdrawn from consideration.)

Claims 34, and 36-39 are pending and are currently under examination.

Priority

A review of the parent applications (08/828,683; 08/625,328; 08/710,802) revealed priority to application No. 08/710,802 filed 09/23/1996. If applicant disagrees with any rejection of claims 34, 36-39 set forth in this office action based on examiner's establishment of a priority date of **September 23, 1996** for the instant claims in application serial number 09/993,234, applicant is invited to submit evidence pointing to the serial number, page and line where support can be found establishing an earlier priority date.

Art Unit: 1642

Information Disclosure Statement

References 210-233, and 235 are improperly cited and will not be printed for publication because they lack publication dates. See 37 CFR 1.98 (b) (5).

Specification

The attempt to update the priority status of the present application as set forth in the preliminary amendment of Paper No. 6 is incomplete. Applicants should indicate that U.S. Serial no. 08/828,683 filed March 31, 1997 is now U.S. Patent No. 6,469,144.

The description of the figures on page 11 (in particular Figures 1-2, 3, 5-6) in which sequences are referenced is incomplete for improper disclosure of amino acid sequences without inclusion of a sequence identifier (i.e., SEQ ID NO:). Thus, this application fails to comply with the requirements of 37 CFR 1.821 through 1.825. This definition sets forth limits, in terms of numbers of amino acids and/or numbers of nucleotides, at or above which compliance with the sequence rules is required. Amino acid sequences as used in 37 CFR 1.821 through 1.825 are interpreted to mean an unbranched sequence of four or more amino acids.

Thus, applicant should amend the specification to insert the appropriate sequence identifier of each referred to sequence. If these sequences have *not* been previously disclosed in a CRF, applicant must provide a computer readable form (CRF) of the sequence listing, an initial or substitute paper copy of the sequence listing, as well as any amendment directing its entry into the specification, and a statement that the content of the paper and computer readable

Art Unit: 1642

copies are the same and, where applicable, include no new matter, as required by 37 CFR 1.821(e-f) or 1.825(b) or 1.825(d). Failure to supply the appropriate sequences identification numbers in response to this action will be considered non-responsive.

Claim Rejections - 35 USC § 112

The following is a quotation of the second paragraph of 35 U.S.C. 112:

The specification shall conclude with one or more claims particularly pointing out and distinctly claiming the subject matter which the applicant regards as his invention.

Claim 39 is rejected under 35 U.S.C. 112, second paragraph, as being incomplete for omitting essential steps, such omission amounting to a gap between the steps. See MPEP § 2172.01. The claim is drawn to a process of using a nucleic acid molecule encoding Apo-3 to effect production of Apo-3. It is not distinctly clear how using the nucleic acid will "effect production". Thus, there appears to be a step missing. The rejection can be obviated by amending the claim to a process of producing an Apo-3 polypeptide comprising culturing the host cell of Claim 38 and isolating said polypeptide.

The following is a quotation of the first paragraph of 35 U.S.C. 112:

The specification shall contain a written description of the invention, and of the manner and process of making and using it, in such full, clear, concise, and exact terms as to enable any person skilled in the art to which it pertains, or with which it is most nearly connected, to make and use the same and shall set forth the best mode contemplated by the inventor of carrying out his invention.

Claim 34 is rejected under 35 USC 112, first paragraph, as the specification does not contain a written description of the claimed invention. The limitation of isolated nucleic acid

Art Unit: 1642

encoding Apo-3 polypeptide comprising amino acid residues 25-417 has no clear support in the specification and the claims as originally filed. Hence, this is a new matter rejection. Applicant should cancel this terminology or submit evidence pointing to the serial number, page and line where support can be found for the disputed terminology.

Claims 34, and 36-39 are rejected under 35 U.S.C. 112, first paragraph, as containing subject matter which was not described in the specification in such a way as to reasonably convey to one skilled in the relevant art that the inventor(s), at the time the application was filed, had possession of the claimed invention.

The written description in this case only sets forth an isolated nucleic acid encoding Apo-3 polypeptide comprising amino acid residues 1 to 417, 25 to 198, or 338 to 417 of SEQ ID NO:6. Therefore the written description is not commensurate in scope with the claims inclusive of isolated nucleic acids encoding "a biologically active variant thereof".

To provide adequate written description and evidence of possession of a claimed genus, the specification must provide sufficient distinguishing identifying characteristics of the genus. The factors to be considered include disclosure of complete or partial structure, physical and/or chemical properties, functional characteristics, structure/function correlation, methods of making the claimed product, or any combination thereof. In this case, no such factors are attributed to the claimed variant. Further, there is no identification of any particular portion of the variant that must be conserved. Accordingly, in the absence of sufficient recitation of distinguishing identifying characteristics, the specification does not provide adequate written description of the claimed genus.

Art Unit: 1642

Vas-Cath Inc. v. Mahurkar, 19USPQ2d 1111, clearly states "applicant must convey with reasonable clarity to those skilled in the art that, as of the filing date sought, he or she was in possession of the invention. The invention is, for purposes of the 'written description' inquiry, whatever is now claimed." (See page 1117.) The specification does not "clearly allow persons of ordinary skill in the art to recognize that [he or she] invented what is claimed." (See Vas-Cath at page 1116). As discussed above, the skilled artisan cannot envision the detailed chemical structure of nucleic acids encoding the encompassed genus of variant polypeptides, and therefore conception is not achieved until reduction to practice has occurred, regardless of the complexity or simplicity of the method of isolation. Adequate written description requires more than a mere statement that it is part of the invention and reference to a potential method of isolating it. The compound itself is required. See Fiers v. Revel, 25 USPQ2d 1601 at 1606 (CAFC 1993) and Amgen Inc. v. Chugai Pharmaceutical Co. Ltd., 18 USPQ2d 1016.

One cannot describe what one has not conceived. See *Fiddes v. Baird*, 30 USPQ2d 1481 at 1483. In *Fiddes*, claims directed to mammalian FGF's were found to be unpatentable due to lack of written description for that broad class. The specification provided only the bovine sequence.

Therefore, only isolated nucleic acid encoding Apo-3 polypeptide comprising amino acid residues 1 to 417, 25 to 198, or 338 to 417 of SEQ ID NO:6, but not the full breadth of the claim meets the written description provision of 35 U.S.C. §112, first paragraph. Applicant is reminded that *Vas-Cath* makes clear that the written description provision of 35 U.S.C. §112 is severable from its enablement provision (see page 1115).

Art Unit: 1642

Claim Rejections - 35 USC § 102

The following is a quotation of the appropriate paragraphs of 35 U.S.C. 102 that form the basis for the rejections under this section made in this Office action:

A person shall be entitled to a patent unless -

(e) the invention was described in (1) an application for patent, published under section 122(b), by another filed in the United States before the invention by the applicant for patent or (2) a patent granted on an application for patent by another filed in the United States before the invention by the applicant for patent, except that an international application filed under the treaty defined in section 351(a) shall have the effects for purposes of this subsection of an application filed in the United States only if the international application designated the United States and was published under Article 21(2) of such treaty in the English language.

Claims 34, 36-39 are rejected under 35 U.S.C. 102(e) as being anticipated by Yu et al. (US Patent No. 6,153,402; earliest priority date is March 12, 1996; IDS #25 and #26).

Yu et al. teach an isolated nucleic acid encoding amino acid residues 25-198 (see attached sequence comparison; also see page 67 of US Provisional 60/013,285 beginning at amino acid position No. 36). Yu et al. further teach vectors, host cells, and a process of using said host cells to effect production of the polypeptide comprising culturing the host cell (page 5, 2nd paragraph, of 60/013,285).

Claims 34, and 36-39 are also rejected under 35 U.S.C. 102(e) as being anticipated by Feldmann et al. (US Patent No. 5633145; earliest priority date is May 30, 1993)

Feldmann *et al.* teach an isolated nucleic acid encoding a biologically active variant polypeptide. (see attached sequence comparison; isolated DNA encoding the variant polypeptide is 39% similar to SEQ ID NO:6). Feldmann *et al.* further teach recombinant expression of biologically active variant polypeptides via host cells transfected with vectors (column 5).

No claim is allowed.

Any inquiry concerning this communication or earlier communications from the examiner should be directed to Gary B. Nickol Ph.D. whose telephone number is 703-305-7143. The examiner can normally be reached on M-F, 8:30-5:00 P.M..

If attempts to reach the examiner by telephone are unsuccessful, the examiner's supervisor, Anthony Caputa can be reached on 703-308-3995. The fax phone numbers for the organization where this application or proceeding is assigned are 703-305-3014 for regular communications and 703-308-4242 for After Final communications.

Any inquiry of a general nature or relating to the status of this application or proceeding should be directed to the receptionist whose telephone number is 703-308-0196.

Gary B. Nickol, Ph.D. Examiner Art Unit 1642

GBN

October 3, 2003

Pary Bricker

옻

Run

```
Title:
Perfect so
Sequence:
                                                                                                                                                                                                                                                                                                 Command line parameters:
-MODEL=frame+_p2n.model -DEV=xlh
-Q-/cgn2_1/USPTO_spool/US09993234/runat_27032003_115457_15378/app_query.fasta_
-Q-/cgn2_1/USPTO_spool/US09993234/runat_27032003_115457_15378/app_query.fasta_
-DB=Issued_Patents_NA -QFMT=fastap -SUFFIX=rni -MINMATCH=0.1 -LOOPCL=0
-LOOPEXT=0 -UNITS=bits -START=1 -END=-1 -MATRIX=blosum62 -TRANS=human40.cdi
-LIST=45 -DOCALIGN=200 -THR_SCORE=pct -THR_MAX=100 -THR_MIN=0 -ALIGN=15
-MODE=LOCAL -OUTFMT=pto -NORM=ext -HEAPSIZE=500 -MINLEN=0 -MAXLEN=200000000
-USER=US0993234_@CGN_1_116_@runat_27032003_115457_15378 -NCPU=6 -ICPU=3
-NO_XLPXY -NO_MMAP -LARGEQUERY -NEG_SCORES=0 -WAIT -LONGLOG -DEV_TIMEOUT=120
-WARN_TIMEOUT=30 -THREADS=1 -XGAPOP=10 -XGAPEXT=0.5 -FGAPOP=6 -FGAPEXT=7
-YGAPOP=10 -YGAPEXT=0.5 -DELOP=6 -DELEXT=7
                                                                                                                                                                                                                                                                                                                                                                                                                                                            Maximum
                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Minimum
                                                                                                                                         Result
                                                                                                                                                                                                                                                                                Database
                                                                                                                                                                                                                                                                                                                                                                                                                                          Post-processing:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             Searched:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      Scoring table:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       protein
                                                                                                                                ŏ
                                                                                                                                                                                Pred.
score
and is
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          number
                                                                                                                                                                                                                                                                                                                                                                                                                                                              DB
BC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 score:
 1038
1038
1038
1038
950
950
666
212.5
212.5
211.5
                                                                                                                                                                                  SŢ
                                                                                                                                core
                                                                                                                                                                                                                                                                                                                                                                                                                                                             seq
                                                                                                                                                                                                                                                                                                                                                                                                                                                                     seq
                                                                                                                                                                                                    o
.
                                                                                                                                                                                          greater
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         o<sub>f</sub>
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       nucleic search,
                                                                                                                                                                                  derived
                                                                                                                                                                                                                                                                                                                                                                                                                                                             length:
length:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        US-09-993-234-6_0
1038
1 QGGTPerra
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              Xgapop
Ygapop
Fgapop
Delop
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          hits
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    April
                                                                                                                              %
Query
Match
                                                                                                                                                                                                                    Issued_Patents_NA:*
1: /cgn2_6/ptodata/1/ina/5A_COMB.seq:*
2: /cgn2_6/ptodata/1/ina/5B_COMB.seq:*
3: /cgn2_6/ptodata/1/ina/6A_COMB.seq:*
4: /cgn2_6/ptodata/1/ina/6B_COMB.seq:*
5: /cgn2_6/ptodata/1/ina/PCTUS_COMB.seq:*
6: /cgn2_6/ptodata/1/ina/backfiles1.seq:*
                                                                                         100.0
                                                                                                                                                                                                   18
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             441362
                                                                              100.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       BLOSUM62
                                                                                                                                                                                                                                                                                                                                                                                                                                         Minimum
                                                                                                                                                                                                                                                                                                                                                                                                                      Listing
                                                                                                                                                                                                                                                                                                                                                                                                                                Maximum
                              91.
64.
20.
   20.
20.
19.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         QGGTRSPRCDCAGDFHKKIG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Copyright
                                                                                                                                                                                                   the
                                                                                                                                                                                         than
    845500550
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        satisfying chosen
                                                                                                                                                                                γd
                                                                                                                                Length
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               10.0
10.0
6.0
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    6,
                                                                                                                                                                                                  number of
                                                                                                                                                                                                                                                                                                                                                                                                                                                             2000000000
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             segs,
                                                                                                                                                                              n or equal analysis c
                                                                                                                                                                                                                                                                                                                                                                                                                       Match 0%
Match 100
first 45
  1254
1634
1783
1783
1438
1438
1438
1049
1049
1956
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   2003,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   GenCore
(c) 1993
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      using
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             153338381
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               Xgapext
Ygapext
Fgapext
Delext
                                                                                                                                                                                                                                                                                                                                                                                                                       100%
45 SI
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            _COPY_25
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    16:28:24
US-08-815-469-3
US-08-928-069-11
US-08-828-683A-9
US-08-815-469-1
US-08-928-069-5
US-08-828-683A-5
US-08-928-069-2
US-08-828-683A-2
US-08-804-166-1
US-08-910-991-1
US-08-762-308-10
US-08-804-166-7
                                                                                                                                                                                         results
                                                                                                                                                                                of to
                                                                                                                                                                                                                                                                                                                                                                                                                       summaries
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      frame_plus_p2n
                                                                                                                                                                                the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    version - 2003
                                                                                                                                                             SUMMARIES
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         parameters
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             residues
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   7700
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 . . . . . . . . . . . . .
                                                                                                                                                                                total
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             .98
                                                                                                                                                                                          predicted by score of the
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                (without 1816.682
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Search time
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   5.1.4_p5_
Compugen
                                                                                                                                                                                score
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          CPTSTLGSCPERCAAVCGWR
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      model
                                                                                                                                                                                          the
                                                                                                                                                                               by chance to he he result being distribution.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                alignments)
Million cell
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    _4578
Ltd.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    29.3732
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          882724
                                                                                                                               Description
        Sequence
                                                                                                                                                                                        to hav
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Seconds
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               updates/sec
                                                                                                                                                                                         printed,
                                                                                                                                                                                                    ρ
           Ó
                              Appli
Appli
Appli
Appli
Appli
Appli
Appli
Appli
Appli
 Appli
Appli
                    Appli
                                                                                                                                                                                                                                                                                                                                                                                 2346
                                                                                                                                                                                                                                                                                                                                       RESULT 1
US-08-815-469-3
                                                                                                                                                                                                                                                                                                                                                                                                                                444033554311144
44403355431144
44133754331355431144
                                                                                                                                                                                                                      ADDRE
                                                                                                                                                                                                                      SSEE:
```

| ភ | 4 | ω | N | 1 | 0 | 9 | 80 | 7 | σv | 5 | 4 | ω | N | - | 0 | 9 | œ | 7 | σ | Ņ | 4 | w | Ŋ | _ | 0 | φ | œ | 7 | ס | u | 4 | ω |
|-----------------|-----------------|-----------------|-------------------|-------------------|-----------------|-----------------|-----------------|-------------------|----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-----------------|-------------------|-----------------|
| 70. | | 16 | $\mathbf{\sigma}$ | $\mathbf{\sigma}$ | ന | m | \mathbf{c} | $\mathbf{\alpha}$ | B | 2 | S | S | Ö. | 00. | \circ | \circ | \circ | 0 | C | 0 | \circ | 0 | 0 | 0 | 0 | 0 | 0 | ω | 03. | 0 | 205 | • |
| 6 | 6 | 7. | 7 | 7 | 7 | 7. | 7. | 8 | 8 | 8 | æ | 8 | 9. | 9 | 9. | 9. | 9. | 9. | 9. | 9 | 9. | 9 | 9 | 9. | 9 | 9 | 9 | 9 | 9. | 9 | 19.7 | |
| ~1 | ~1 | 62 | A 3 | 92 | 59 | 55 | 57 | 72 | 72 | 72 | 44 | 54 | 47 | 48 | 14 | 14 | 89 | 88 | 88 | 17 | \neg | 17 | 17 | 16 | 16 | 9 | 90 | 20 | 20 | 0 | 600 | 0 |
| N | ب | 4 | 4 | 4 | 4 | 4 | 4 | 4 | 4 | Çī | 4 | 4 | 4 | 4 | 4 | 4 | N | σ | μ. | 4 | | _ | - - | 4 | W | N | 1 | 4 | 4 | N | – | 4 |
| S-08-465-982-5 | S-08-050-319E | S-09-333-279-6 | S-08-509-024- | S-09-146-950- | S-09-146-950- | S-09-146-950-1 | S-09-146-950- | S-09-333-279- | S-08-509-024- | CT-US96-12374 | S-09-513-007- | S-09-513-007- | S-09-149-922- | S-09-326-394- | S-08-910-991- | S-08-804-166- | S-08-627-151A- | CT-US95-09576 | S-08-286-740- | S-08-054-970- | S-08-126-016- | S-08-837-941- | S-08-321-668- | S-09-505-250- | S-09-106-038A- | S-08-465-982- | S-08-050-319B- | S-08-910-991- | S-08-804-166- | S-08-465-982-4 | US-08-050-319B-47 | S-08-910-991- |
| equence 53, App | equence 53, App | equence 6, Appl | equence 6, Appl | equence 1, Appl | equence 17, App | equence 19, App | equence 3, Appl | equence 1, Appl | equence 1, App | equence 1, Appl | equence 1, Appl | equence 3, Appl | equence 6, Appl | equence 1, Appl | equence 5, Appl | equence 5, Appl | equence 6, Appl | equence 2, Appl | equence 2, Appl | equence 1, Appl | equence 1, App | equence 1, Appl | equence 1, Appl | equence 3, Appl | equence 1, Appl | equence 24, App | equence 24, App | equence 3, Appl | equence 3, Appl | equence 47, App | ָׁ <mark>ַ</mark> | equence 7, Appl |

ALIGNMENTS

```
Sequence 3, Application US/08815469

Patent No. 6153402

GENERAL INFORMATION:
APPLICANT: Yu, Guo-Liang
APPLICANT: Ni, Jian
APPLICANT: Dixit, Vishva
APPLICANT: Gentz, Reiner L.
APPLICANT: Dillon, Patrick J.
TITLE OF INVENTION: Death Domain Containing Receptors
NUMBER OF SEQUENCES: 17
CORRESPONDENCE ADDRESS:
          CLASSIFICATION: 435
PRIOR APPLICATION DATA:
APPLICATION NUMBER: No. 61
FILING DATE: 06-FEB-1997
PRIOR APPLICATION DATA:
APPLICATION UMBER: US 60/FILING DATE: 17-00"
                                                                                                                                             STREET: 1100 New York Ave., NW, Suite 600
CITY: Washington
STATE: DC
COUNTRY: USA
ZIP: 20005-3934
COMPUTER READABLE FORM:
MEDIUM TYPE: Floppy disk
COMPUTER: IBM PC compatible
OPERATING SYSTEM: PC-DOS/MS-DOS
SOFTWARE: Patentin Release #1.0, Version #1
CURRENT APPLICATION DATA:
APPLICATION NUMBER: US/08/815,469
FILING DATE: HEREWITH
CLASSIFICATION: 435
US 60/028,711
PPLICATION DATA:
CATION NUMPER
                                                                                                                                                                                                                                                                                                                                                                                     1100 New York
                                                                                                                                                                                                                                                                                                                                                                                     Kessler,
ork Ave.,
                                                                                                                 6153402 Yet
                                                                                                                                                                                                                                                                                                                                                                                   Goldstein
NW, Suite
                                                                                                                Assigned
                                                                                                                                                                                                                                                                                                                                                                                     & Fox,
                                                                                                                                                                                                                          #1.30
                                                                                                                                                                                                                                                                                                                                                                                                       P.L.L.C
```

```
Score:
Percent Similarity:
Best Local Similarity:
Query Match:
DB:
                                                                                           δÃ
                                                                 Ş
                                                                               멍
                                                                                                         밁
                                                                                                                                                                                                                                                                                Š
                                  US-08-928-069-11
                                        RESULT
                                                                                                                      Ş
                                                                                                                                                                                                                            Ã
                                                                                                                                                                                                                                           B
                                                                                                                                                                                                                                                       Š
                                                                                                                                                                                                                                                                     9
                                                                                                                                                                                                                                                                                             US-09-993-234-6_COPY_25_198
                                                                                                                                                                                                   Š
                                                                                                                                                                                                                 g
                                                                                                                                   9
                                                                                                                                                                                       9
                                                                                                                                                                          Š
                                                                                                                                                                                                                                                                                                                                                                                                              REGISTRATION NUMBER: 36,688
REFERENCE/DOCKET NUMBER: 1488
TELECOMMUNICATION INFORMATION:
TELEPHONE: 202-371-2600
TELEFAX: 202-371-2540
INFORMATION FOR SEQ ID NO: 3:
SEQUENCE CHARACTERISTICS:
LENGTH: 1254 base pairs
         Sequence 11, Applicate Patent No. 6462176
GENERAL INFORMATION:
                                                                                                                                                                                                                                                                                                                                                              LOCATION: 08-815-469-3
                                                                                                                                                                                                                                                                                                                                                  nment
                                                                                                                                                                                                                                                                                                                                                                                    MOLECULE TYPE: FEATURE:
 APPLICANT:
                                                                                                                                                                                                                                                                                                                                                                                   FEATURE
                                                      553
                                                                                                          433
                                                                                                                                                                                                                                                                                                                                            NO.
                                                                                                                                                                                                                                                                                                                                                                                              STRANDEDNESS: OTOPOLOGY: both
                                                                                                                                                                                                                                           133
                                                                                                                                                101
                                                                                                                                                                                                                 193
                                                                                                                                                             313
                                                                                                                                                                                       253
                                                                                                                                                                                                                             41
                                                                                                                                                                                                                                                                    73
                                                                                                                                                                                                                                                                                                                                                                            NAME/KEY:
                                                                                                                                                                                                                                                       21
                                                                                                                                                                          48
                                                                                                                                                                                                   61
                                                                                                                                                                                                                                                                                ۳
                                                   GlySerCysProGluArgCysAlaAlaValCysGlyTrpArg
                                                                                                       11,
6462
                                                                             Scores:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    DATE
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Steffe,
: Ashkenazi, INVENTION: A
                     Application
162176
                                                                                                                                                                                                                                                                                                                                                                    CDS
1..
                                                                                                                                                                                                                                                                                                                                                                                                both
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            12-MAR-1996
INFORMATION:
                                                                                                                                                                                                                                                                                                                                                                                         CDNA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Eric
                                                                                                                                                                                                                                                                                                                                                                                                            acid
                                                                                                                                                                                                                                                                                                                                                                                                      double
                                                                                                                                                                                                                                                                                                          3.22e-86
1038.00
100.00%
100.00%
100.00%
3
l, Avi
Apo-3
                                                                                                                                                                                                                                                                                                                                                                                                                                                                       ⊼
                           US/08928069
                                                                                                                                                                                                                                                                                             (1-174)
                                                                                                                                                                                                                                                                                                                                                                                                                                                          36,688
R: 1488.0310003/EKS/KRM
J.
POLYPEPTIDE
                                                                                                                                                                                                                                                                                             ×
                                                                                                                                                                                                                                                                                             US-08-815-469-3
                                                                                                                                                                                                                                                                                                        Length:
Matches:
Conservative:
Mismatches:
Indels:
Gaps:
                                                                                                                                                                                                                                                                                                                                    1254
174
                                                                                                                                                                                                                                                                                             (1-1254)
                                                                                                                                                                                                                                                                                                           0000
                                                                                                                    140
                                                                              55
                                                                                           160
                                                                                                        492
                                                                                                                                  432
                                                                                                                                              120
                                                                                                                                                            372
                                                                                                                                                                        100
                                                                                                                                                                                                  80
                                                                                                                                                                                                                            60
                                                                                                                                                                                                                                         192
                                                                                                                                                                                                                                                                   132
                                                                                                                                                                                     312
                                                                                                                                                                                                                                                      40
                                                                                                                                                                                                                                                                                20
```

0

```
Qy
                                                                                                                                                                                                                                                                                                                   Score:
                     δŌ
                                        멍
                                                         Ş
                                                                            밁
                                                                                                                맔
                                                                                                                                 Ş
                                                                                                                                                    B
                                                                                                                                                                                         В
                                                                                                                                                                                                          Ş
                                                                                                                                                                                                                            밁
                                                                                                                                                                                                                                            Ş
                                                                                                                                                                                                                                                              US-09-993-234-6.
                                                                                                                                                                                                                                                                                  DB:
                                                                                            γQ
                                                                                                                                                                                                                                                                                                    Percent Sin
Best Local
                                                                                                                                                                                                                                                                                          Query
                                                                                                                                                                                                                                                                                                                                      Alignment
                                                                                                                                                                                                                                                                                                                               Pred.
                                                                                                                                                                                                                                                                                                                                                                                        NAME: Marschang, Diane L.
REGISTRATION NUMBER: 35,600
REFERENCE/DOCKET NUMBER: P105
TELECOMMUNICATION INFORMATION:
TELEPHONE: 650/225-5416
TELEPHONE: 650/952-9881
INFORMATION FOR SEQ ID NO: 11:
SEQUENCE CHARACTERISTICS:
LENGTH: 1634 base pairs
                                                                                                                                                                                                                                                                                                                                                       LENGTH: 1634
TYPE: Nucleic
STRANDEDNESS:
TOPOLOGY: Lin
-08-928-069-11
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 SOFTWARE: WinPatin (Gene
CURRENT APPLICATION DATA:
APPLICATION NUMBER: US/(
FILING DATE: 11-Sep-1997
CLASSIFICATION: 435
                                                                                                                                                                                                                                                                                                                                                                                                                                                                    FILING DATE: 09/23/1996
ATTORNEY/AGENT INFORMATION:
NAME: Marschang, Diane L
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               COMPUTER READABLE FORM:
MEDIUM TYPE: 3.5 inc
COMPUTER: IBM PC com
OPERATING SYSTEM: PC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          NUMBER OF SEQUENCES: 1
CORRESPONDENCE ADDRESS:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                PRIOR APPLICATION DATA:
APPLICATION NUMBER:
                                                                                                                                                                                                                                                                                                                               No.:
                                                                                                                                                                                                                                                                                           Match:
                                        461
                                                         101
                                                                           401
                                                                                                                341
                                                                                                                                                                                         221
                                                                                                                                                    281
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      STREET:
CITY: :
STATE:
                                                                                                                                                                                                                            61
                                                                                             81
                                                                                                                                                                      41
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   ZIP:
                                                                                                                                  61
                                                                                                                                                                                                                                                                                                  Similarit cal Simila
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    COUNTRY: US
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  ADDRESSEE
                                    Scores
                                                                                                                                                                                                                                                                                                                                                                  Nucleic Acid
)EDNESS: Single
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               South San
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               California
: USA
                                                                                                                                                                                                                                                            _COPY_25_198 (1-174) x US-08-928-069-11
                                                                                                                                                                                                                                                                                                 arity:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1 DNA Way
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             IBM PC compatible SYSTEM: PC-DOS/MS-DOS
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       WinPatin (Genentech)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Genentech,
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          лывк: US/08/928,069
11-Sep-1997
                                                                                                                                                                                                                                                                                4.57e-86
1038.00
100.00%
100.00%
100.00%
100.00%
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Francisco
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                60/026943
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  Inc.
                                                                                                                                                                                                                                                                                                                                                                                                                                                    P1052R1
                                                                                                                                                                                                                                                                                                 Conservative: Mismatches:
                                                                                                                                                                                                                                                                                Gaps:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  floppy disk
                                                                                                                                                                                                                                                                                          Indels:
                                                                                                                                                                                                                                                                                                                    Matches:
                                                                                                                                                                                                                                                                                       1634
174
0
                                                                                                                                                                                                                                                             (1-1634)
```

100

400

08

60

280

40

220

20

340

580

140

520

120

460

(1-417) x US-08-050-319B-24 (1-2062)

```
8
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       Ş
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         멍
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           Ş
   Query
                                            Pred
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Ş
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    20
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            B
                    Best
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   B
                           Percent Similarity
                                      S
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             5
                                                      Alignment
                                      core:
                                                                                                                                                                                                                                                                                                                                                                                                                                  Patent No.
                                                                                                                                                    TELECOMMUNICATION INFORMATION:
TELEPHONE: (415) 617-8999
TELEFAX: (415) 327-3231
INFORMATION FOR SEQ ID NO: 24:
SEQUENCE CHARACTERISTICS:
                                                                         08-050-
                                                                                                                                                                                                                                                                                                                                                                                                                            GENERAL
                                                                                                                                                                                                                CURRENT APPLICATION DATA:
APPLICATION NUMBER: US/08,
FILING DATE: 10-May-1993
CLASSIFICATION: 435
ATTORNEY/AGENT INFORMATION:
NAME: Robbins, Roberta L.
           Local S. y Match:
                                                                                                                                                                                                                                                                                                                                                                                                                                                     8-050-319B-
                                                                                                                                                                                                                                                                                                                                                                                       APPLICANT: M.J.C. Turner, F.M Brennan
TITLE OF INVENTION: Modified human TNFalpha
TITLE OF INVENTION: Necrosis Factor alpha)
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1653
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         1473
                                                                                                                                                                                                                                                                                                 COMPUTER READABLE FORM: MEDIUM TYPE: Floppy
                                                                                                                                                                                                                                                                                                                                                                      NUMBER OF SEQUENCES: 5
CORRESPONDENCE ADDRESS:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   1593
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      1533
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             1413
                                                                                                         MOLECULE TYPE:
                                                                                                                                                                                                                                                                                                                                                                                                                 APPLICANT:
                                                                                                  FEATURE
                                              NO:..
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           354
                                                                                                                        TYPE: nuclei
                                                                                                                                                                                              NAME: Robbins, Roberta REGISTRATION NUMBER: 33 REFERENCE/DOCKET NUMBER:
                                                                                                                                                                                                                                                                                                                                                             ADDRESSEE:
                                                                       LOCATION: 50-319B-24
                                                                                        NAME/KEY:
                                                                                                                                                                                                                                                                                OPERATING SYSTEM:
                                                                                                                    TOPOLOGY:
                                                                                                                                                                                                                                                                       SOFTWARE:
                                                                                                                                                                                                                                                                                         COMPUTER:
                                                                                                                                                                                                                                                                                                                             COUNTRY:
                                                                                                                                             LENGTH: 2062 base
                                                                                                                                                                                                                                                                                                                                             STREET:
                                                                                                                                                                                                                                                                                                                   IP:
                                                                                                                                                                                                                                                                                                                                     TATE:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     CTGGGACTGAGCGAGCACGAGATCGAGCGGCTGGAGCTGGAGAACGGGCGCCACCTGCGC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ACCCTGTACGCGGTGGTGGACGGCGTGCCCCCGTCGCGCTGGAAGGAGTTGGTGCGGCGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                               GAGGAGGCGCTG 1664
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 CTGGAGCTGCTGGGCCGCGTGCTCAGGGACATGGACCTGCTGGGTTGCCTGGAAAACATA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                LeuGlyLeuArgGluAlaGluIleGluAlaValGluValGluIleGlyArg-
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    GlnLeuTyrAspValMetAspAlaValProAlaArgArgTrpLysGluPheValArgThr
                                                       Scores
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ArgSerArgLeu
                 "LIGITICY:
Similarity:
                                                                                                                                                                                                                                                                                                                                                                                                                           INFORMATION:
                                                                                                                                                                                                                                                                                                                                                                                                                                          24
                                                                                                                                                                                                                                                                                                                                   T: 635 Bryant:
Palo Alto
California
                                                                                                                                                                                                                                                                                                                                                                                                                                  4, Application 5633145
                                                                                                                                                                                                                                                                                                                   94301
                                                                                                                                    nucleic
                                                                                                                                                                                                                                                                                                                           USA
                                                                                                                                                                                                                                                                                                                                                                                                                  M. Feldmann,
                                                                                CDS
155.
                                                                                                                                                                                                                                                                       PatentIn Release
                                                                                                                                                                                                                                                                                         IBM PC
                                                                                                                   linear
                                                                                                                                                                                                                                                                                                                                                              Reed & Robbins
                                                                                                                                                                                                                                       лмькк: US/08/050,319B
10-May-1993
                                                                                                                                                                                                                                                                                                Floppy disk
                                                                                                          CDNA to
                                                                                                                                   acid
                                                                                                                            double
                                                                                 .1519
         1.36e-21
376.00
39.09%
28.29%
16.19%
                                                                                                                                                                                                                                                                               PC-DOS/MS-DOS
                                                                                                                                           pairs
                                                                                                                                                                                                                                                                                                                                                                                                                                           US/08050319B
                                                                                                                                                                                                                                                                                                                                                      Street
                                                                                                                                                                                                                                                                                                                                                                                                                 P.W.
                                                                                                                                                                                                      33,208
                                                                                                           mRNA
                                                                                                                                                                                                 5150-0030
                                                                                                                                                                                                                                                                                                                                                                                                                  Gray,
                                                                                                                                                                                                                                                                       #1.0,
Mismatches:
Indels:
Gaps:
                          Matches:
Conservati
                                             Length:
                                                                                                                                                                                                                                                                       version
                          ive:
                                                                                                                                                                                                                                                                                                                                                                                       Receptor
                                                                                                                                                                                                                                                                                                                                                                                               (Tumor
2062
131
50
182
100
22
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        -PheArg
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      AlaGly
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           1472
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  165
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1592
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        1532
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    409
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             353
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          372
                                                                                                                 Š
                                                                                                                                                                                                                                                                                                QΥ
                                                                                                                                                                                                                                                                                                                                                                                                       Qy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Ş
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    Ş
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      멼
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       γQ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          γQ
                                                                                                                                   밁
                                                                                                                                                                                                                                                                                                                                                                     Ş
                                                                                                                                                                                                                                                                                                                                                                                                                                          Ş
                                                                                                                                                                                                                                                                                                                                                                                                                                                                            Š
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         멍
        Ş
                           밁
                                           δÃ
                                                             g
                                                                              S
                                                                                                Db
                                                                                                                                                   Ş
                                                                                                                                                                     뫄
                                                                                                                                                                                      Ş
                                                                                                                                                                                                        8
                                                                                                                                                                                                                        QΥ
                                                                                                                                                                                                                                            멍
                                                                                                                                                                                                                                                            Ş
                                                                                                                                                                                                                                                                               밁
                                                                                                                                                                                                                                                                                                                  g
                                                                                                                                                                                                                                                                                                                                   Š
                                                                                                                                                                                                                                                                                                                                                    D
D
                                                                                                                                                                                                                                                                                                                                                                                       B
                                                                                                                                                                                                                                                                                                                                                                                                                          g
                                                                                                                                                                                                                                                                                                                                                                                                                                                             밁
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                멍
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            -09-993-234-6
                                                             1037
                                                                                                                  25
                          064
                                           293
                                                                                                977
                                                                                                                                  926
                                                                                                                                                                     869
                                                                                                                                                                                                        809
                                                                                                                                                                                                                                                             196
                                                                                                                                                                                                                                                                              701
                                                                                                                                                                                                                                                                                                181
                                                                                                                                                                                                                                                                                                                 644
                                                                                                                                                                                                                                                                                                                                   161
                                                                                                                                                                                                                                                                                                                                                    599
                                                                                                                                                                                                                                                                                                                                                                      141
                                                                                                                                                                                                                                                                                                                                                                                       554
                                                                                                                                                                                                                                                                                                                                                                                                        124
                                                                                                                                                                                                                                                                                                                                                                                                                          494
                                                                                                                                                                                                                                                                                                                                                                                                                                           104
                                                                                                                                                                                                                                                                                                                                                                                                                                                             437
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   317
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      257
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         197
                                                                                                                                                    240
                                                                                                                                                                                       227
                                                                                                                                                                                                                          207
                                                                                                                                                                                                                                                                                                                                                                                                                                                                             84
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 64
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    44
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          15
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       27
                                                             TCC-
```

```
1109
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         TGCCTC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              CysLeu
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  GlyMet
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       GGGGAC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               CICCIG
    CAGGGG
                                                                                                            Cyspro
                                                                                                                                                                                     Thrval
                                                                                                                                                                                                                                                                                               GAGCTT
                                                                                                                                                                                                                                                                                                                                                                                                        CysTrp
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    AspCys
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 CATTAT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    GluCys
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   HisHis
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       CCGGGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          ProCys
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  GlyLeu
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           GlyThr
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  LeuLeu
                                                                         TGTCCC
                                                                                                                                                                                                                         CCCACT
                                                                                                                                                                                                                                                                                                                                                                                                                                              TTCTTT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 LeuAla
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ATTGAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 ThrSer
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               CACCIC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              AATTCG.
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ACCGTG
                                                                                                                                                                                                                                                                                                                                                                     TGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          GlnValThrTrpSerTrpAspGlnLeuProSerArgAlaLeuGlyPro----
                                                                                                                                             GlnLeuValGlyAsnSerTrpThrProGlyTyrProGluThrGlnGluAlaLeu
::::::::||||||||
-----AGCTCCACCTATACCCCCGGTGAC-------
                                                                                                                                                                                                                     CCAGGCTTCACCCCACCCTGGGCTTCAGTCCCGTGCCCAGTTCCACCTTCACC
                                                                                                                                                                                                                                                                                           GluAlaLeuThrProProProAlaThrHisLeuSerProLeuAspSer-----
                                                                                                                                                                                                                                                                                                                                                                  AAGTCCAAGCTCTACTCCATTGTTTGTGGGAAATCGACACCTGAAAAAGAGGGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                GlyLeuValValProLeuLeuGlyAlaThrLeuThrTyrThrTyrArgHis
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   TGCACCTGCCATGCAGGTTTCTTTCTAAGAGAAAACGAGTGTGTCTCCTGT----
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    GlyThrCysLeuProGlyPheTyrGluHisGlyAspGlyCysValSerCysPro
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           AAT - - -GGGACCGTGCAC -
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          AspCysGlyAlaLeuHisArgHisThrArgLeuLeuCysSerArgArgAspThr
::: ||| :::||| ||| ||| |||
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              TGGAGTGAAAACCTTTTCCAGTGC------TTCAATTGCAGCCTC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            {\tt AsnSerGluCysAlaArgCysGlnAlaCysAspGluGlnAlaSerGlnValAla}
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   {\tt GlyAsnSerThrCysLeuValCysProGlnAspThrPheLeuAlaTrpGluAsn}
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              PheCysCysArgGlyCysProAlaGlyHisTyrLeuLysAlaProCysThrGlu
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   AGGGAGAAGAGAGATAGTGTGTGTCCCCAAGGAAAATATATCCACCCTCAAAAT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ArgSerProArg------CysAspCysAlaGlyAspPheHisLysLysIle
                                                                          AACTTTGCG-
                                                                                                                                                                                                                                                                                                                                                                                                          ProHisLys---
                                                                                                                                                                                                                                                                                                                                                                                                                                          GGTCTTTGCCTTTTATCCCTCCTCTTCATTGGTTTAATGTATCGCTACCAACGG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 AATGTTAAGGGCACTGAGGACTCAGGCACCACAGTGCTGTTGCCCCTGGTCATT
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             ThrLeuGlySerCysProGlu-----ArgCysAlaAlaValCys-----
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            AGA - - - CACTGCCTCAGCTGCTCCAAATGCCGAAAGGAAATGGGTCAGGTGGAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ATTTGCTGTACCAAGTGCCACAAAGGAACCTACTTGTACAATGACTGTCCAGGC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           ----AGTAACTGTAAGAAAAGCCTGGAGTGCACGAAGTTGTGCCTACCCCAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               --GlnValSerGlnCysValSerSerSerProPheTyrCysGlnPro
                                                                                                                                                                                                                                           -AlaHisThrLeuLeuAlaProProAspSerSerGluLysIleCys
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    ---GlyTrpArgGlnMetPheTrpValGlnValLeu
                                                                         -GCTCCCCGCAGAGAGGTGGCACCACCCTAT
                                                                                                                                                                                                                                                                                                                                                                                                    ----ProLeuValThrAlaAspGluAla
                                     -AlaAlaAlaProThrLeuSerPro---
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         -CTCTCCTGCCAGGAGAAACAGAAC
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         123
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   808
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     206
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           160
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  140
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            376
                                                                                                                                                                                                                                                                                                                                 257
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   700
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      180
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       643
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               195
```

```
δÃ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            DЬ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              QΥ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               В
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         VΩ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   Š
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          B
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           RESULT 12

US-08-465-982-24

US-08-465-982-24

; Sequence 24, Application US/08465982

; Patent No. 5863786

; Patent No. 5863786
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             δõ
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               밁
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        Ş
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   р
; NAME/KEY:
; LOCATION:
US-08-465-982-24
                                                                                                                         NAME: Robbins, Roberta L.
REGISTRATION NUMBER: 33,208
REFERENCE/DOCKET NUMBER: 5150
REFERENCE/DOCKET NUMBER: 5150
RELECOMMUNICATION INFORMATION:
TELEPHONE: (415) 617-8999
TELEFAX: (415) 327-3231
INFORMATION FOR SEQ ID NO: 24:
SEQUENCE CHARACTERISTICS:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               GENERAL INFORMATION:
APPLICANT: M.Feld
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                      1229
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             1169
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                1289
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          1349
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              1469
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   1409
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        319
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   APPLICANT: M.Feldmann, P.W. Gray,
APPLICANT: M.J.C. Turner, F.M Brennan
TITLE OF INVENTION: Modified human TNFalpha (Tumor
TITLE OF INVENTION: Necrosis Factor alpha) Receptor
NUMBER OF SEQUENCES: 57
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 335
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            355
                                                                                                                                                                                                                                                                                                                                   MEDIUM TYPE: Floppy disk
COMPUTER: IBM PC compatible
OPERATING SYSTEM: PC-DOS/MS-DOS
SOFTWARE: PatentIn Release #1.0,
CURRENT APPLICATION DATA:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                             CORRESPONDENCE ADDRESS:
ADDRESSEE: Reed & Ro
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     374
                                                                                                                                                                                                                                                                                                                                                                                            COMPUTER READABLE FORM: MEDIUM TYPE: Floppy
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               391
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          411
                                                                                                                                                                                                                                                                                APPLICATION NUMBER: | FILING DATE: CLASSIFICATION: PRIOR APPLICATION DATA:
                                                                                                                                                                                                                                     APPLICATION NUMBER: US/08/050,319 FILING DATE: 10-May-1993 ATTORNEY/AGENT INFORMATION:
                                                        MOLECULE TYPE:
                                           FEATURE:
                                                                                                                                                                                                                                                                                                                                                                                                                                                   STREET:
CITY:
STATE:
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            ADDRESSEE:
                                                                                                                                                                                                                                                                                                                                                                                                                                        COUNTRY:
                                                                      TOPOLOGY:
                                                                                   STRANDEDNESS
                                                                                                                LENGTH: 2062
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                          CAGAAGTGGGAGGACAGTGCCCACAAGCCACAGAGCCTAGACACTGATGACCCCCGGACG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              GlnGlnTyrGluMetLeuLysArgTrpArgGlnGlnGlnPro---
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  GlyAlaValTyrAlaAlaLeuGluArgMetGlyLeuAspGlyCysValGluAspLeuArg
;;;
GAGCTGCTGGGACGCGTGCTCCGCGACATGGACCTGCTGGGGCTGCCTGGAGGACATCGAG
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         GlyLeuArgGluAlaGluIleGluAlaValGluValGluIleGlyArg---PheArgAsp
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                              GAGGCGCTT 1477
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        SerArgLeu 413
                                                                                                                                                                                                                                                                                                                                                                                                                          94301
                                                                                                                                                                                                                                                                                                                                                                                                                                                      Palo Alto .
California
                                                                                                nucleic acid
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                   635
                                                                                                                                                                                                                                                                                                                                                                                                                                         USA
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                       GluSerProAlaGlySerProAlaMetMetLeuGlnProGlyProGln---
               CDS
155..1519
                                                                      linear
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                Bryant Street
                                                                                                                base pairs
                                                        cDNA to mRNA
                                                                                    double
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 Robbins
                                                                                                                                                                                                                                                                                                                               US/08/465,982
                                                                                                                                                                                                     5150-0030
                                                                                                                                                                                                                                                                                                                                                           version #1.25
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                            -AlaGlyLeu
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                     1228
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                 1288
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  334
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                    373
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                        410
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                         1348
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                           354
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               1468
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                  1408
                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                                               390
```

| 뫄 | γ _Q | Db 43 | ם ב | Qy | dd VQ | QQ VQ | 4d 53 | מם | γQ | Qy da | dg Vg | Qy Db | dd YO' | Оу | ОУ | Оу | Ф | DP QA | US-0 | Alig Pred Scor Perc Best Quer DB: |
|---------------------------------------|-------------------------------------|--|--|--------------------------|---------------------|---------------------------|---|----------------------------|---|--|----------|----------|--|-------------|------------------------|----|---|-------------------------|---|---|
| 1037 TCCAGCTCCACCTATACCCCCGGTGAC 1063 | 273 ThrvalGlnLeuValGlyAsnSerTrpThrP | 977 CCCACTCCAGGCTTCACCCCCACCCTGGGCTTCAGTCCCGTGCC | 926 GAGCTTGAAGGAACTACTACTATEULeuAlaProProAspSerSerGluLysIleCys | 240 GlyMetGluAlaLeuThrPr | 227 CysTrpProHisLys | TGGTTTAATGTATCGCTACCAACGG | GAATGTTAAGGGCACTGAGGACTCAGGCACCACAGTGCT | 01AGTAACTGTAAGAAAAGCCTGGAG | 181 ThrSerThrLeuGlySerCysProGluArgCysAlaAlaValCys 195 | 161 AspCysGlyThrCysLeuProGlyPheTyrGluHisGlyAspGlyCysValSerCysPro 180 | | | ysLysProGIYTIPPHEVAL ::: :: ::: GCAGGAAGAACCAGTACCGG | SASPGluGlni | hrPheLeuAlaTrpGluAsn:: | | | TTATTGGACTGGTCCCTCACCTA | 9-993-234-6 (1-417) x US-08-465-982-24 (1-2062) | Ignment Scores: 1.36e-21 Length: 2062 20. No.: 376.00 Matches: 131 20.09% Conservative: 50 20.29% Mismatches: 182 21.36e-21 Length: 2062 20.09% Matches: 131 20.09% Conservative: 50 20.09% Mismatches: 182 316.19% Indels: 100 32.09% Gaps: 22 |